

<110> CLOUGH; BARBARA PREISER, PETER WILSON, ROBERT

35

<120> AN EF-TU PROTEIN ENCODED ON THE PLASTID DNA OF THE MALARIA PARASITE AND PROTEIN SYNTHESIS INHIBITORS EFFECTIVE AS ANTI-MALARIAL COMPOUNDS

<130> 117-349 <140> US 09/845.335 <141> 2001-05-01 <150> US 09/140.466 <151> 1998-08-26 <150> US 60/056,246 <151> 1997-08-28 <160> 14 <170> Patentin Ver. 2.1 <210> 1 <211> 1230 <212> DNA <213> Plasmodium falciparum <220> <221> CDS <222> (1)..(1230) <400> 1 atg aat aat aaa tta ttt tta aga aat aaa caa cat ata aat tta ggt 48 Met Asn Asn Lys Leu Phe Leu Arg Asn Lys Gln His Ile Asn Leu Gly 1 5 10 15 act ata ggg cat gta gat cat gga aaa act aca tta aca aca gct ata 96 Thr Ile Gly His Val Asp His Gly Lys Thr Thr Leu Thr Thr Ala Ile 20 25 30 tct tat tta tta aat tta caa gga tta tca aaa aaa tat aat tat tca 144 Ser Tyr Leu Leu Asn Leu Gln Gly Leu Ser Lys Lys Tyr Asn Tyr Ser

45

40

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				_		tta Leu 120								384
						aat Asn		-		_	-	_	_	432
			_			tta Leu	_		_					480
						ata Ile								528
						aaa Lys		-		_				576
	Asn					tta Leu 200								624
Asn				Thr		aaa Lys		Asp						672

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a L	ag ys	att Ile	gaa Glu	a caa u G1r	gg 1 G1 24!	y Cy	t at	a aat e Asr	t tt.	a aa u Asi 250	n Ası	t ga p Gl	a att u Ile	ga: G1:	a at [.] u Ile 259	t tta e Leu 5	768
a L	aa ys	ttt Phe	gaa Glu	aaa Lys 260	Sei	a tc Sei	t cct Pro	t aat o Asr	tta Lei 265	ı Thr	a aca Thr	a gti r Va	t ata I Ile	gga G1) 270	/ Let	a gaa u Glu	816
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<212> PRT

<213> Plasmodium falciparum

<400> 2

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Thr Ile Gly His Val Asp His Gly Lys Thr Thr Leu Thr Thr Ala Ile 20 25 30

Ser Tyr Leu Leu Asn Leu Gln Gly Leu Ser Lys Lys Tyr Asn Tyr Ser 35 40 45

Asp Ile Asp Ser Ala Pro Glu Glu Lys Ile Arg Gly Ile Thr Ile Asn 50 55 60

Thr Thr His Ile Glu Tyr Glu Thr Leu Thr Lys His Cys Ala His Ile 65 70 75 80

Asp Cys Pro Gly His Ser Asp Tyr Ile Lys Asn Met Ile Ile Gly Ala 85 90 95

Thr Gln Met Asp Ile Ala Ile Leu Val Ile Ser Ile Ile Asp Gly Ile 100 105 110

Met Pro Gln Thr Tyr Glu His Leu Leu Leu Ile Lys Gln Ile Gly Ile 115 120 125

Lys Asn Ile Ile Ile Phe Leu Asn Lys Glu Asp Leu Cys Asp Asp Val 130 135 140

Glu Leu Ile Asp Phe Ile Lys Leu Glu Val Asn Glu Leu Leu Ile Lys 145 150 155 160

Tyr Asn Phe Asp Leu Asn Tyr Ile His Ile Leu Thr Gly Ser Ala Leu 165 170 175

- Asn Val Ile Asn Ile Ile Gln Lys Asn Lys Asp Tyr Glu Leu Ile Lys 180 185 190
- Ser Asn Ile Trp Ile Gln Lys Leu Asn Asn Leu Ile Gln Ile Ile Asp 195 200 205
- Asn Ile Ile Ile Pro Thr Arg Lys Ile Asn Asp Tyr Phe Leu Met Ser 210 215 220
- Ile Glu Asp Val Phe Ser Ile Thr Gly Arg Gly Thr Val Val Thr Gly 225 230 235 240
- Lys Ile Glu Gln Gly Cys Ile Asn Leu Asn Asp Glu Ile Glu Ile Leu 245 250 255
- Lys Phe Glu Lys Ser Ser Pro Asn Leu Thr Thr Val Ile Gly Leu Glu 260 265 270
- Met Phe Lys Lys Gln Leu Thr Gln Ala Gln Ser Gly Asp Asn Val Gly 275 280 285
- Ile Leu Leu Arg Asn Ile Gln Lys Lys Asp Ile Lys Arg Gly Met Ile 290 295 300
- Leu Ala Thr Pro Asn Lys Leu Lys Val Tyr Lys Ser Phe Ile Ala Glu 305 310 315 320
- Thr Tyr Ile Leu Thr Lys Glu Glu Gly Gly Arg His Lys Pro Phe Asn 325 330 335
- Ile Gly Tyr Lys Pro Gln Phe Phe Ile Arg Thr Val Asp Val Thr Gly 340 345 350
- Glu Ile Lys Asn Ile Tyr Leu Asn Glu Asn Val Gln Lys Val Ala Ile 355 360 365
- Pro Gly Asp Lys Ile Thr Leu His Ile Glu Leu Lys His Tyr Ile Val 370 375 380
- Leu Thr Leu Asn Met Lys Phe Ser Ile Arg Glu Gly Gly Lys Thr Ile 385 390 395 400
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405 410

<210> 3

<211> 409

<212> PRT

<213> Anacystis nidulans

<400> 3

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Thr Thr Val Leu Ala Lys Ala Gly Met Ala Lys Ala Arg Ala Tyr Ala 35 40 45

Asp Ile Asp Ala Ala Pro Glu Glu Lys Ala Arg Gly Ile Thr Ile Asn 50 55 60

Thr Ala His Val Glu Tyr Glu Thr Gly His Arg His Tyr Ala His Val 65 70 75 80

Asp Cys Pro Gly His Ala Asp Tyr Val Lys Asn Met Ile Thr Gly Ala 85 90 95

Ala Gln Met Asp Gly Ala Ile Leu Val Val Ser Ala Ala Asp Gly Pro 100 105 110

Met Pro Gln Thr Arg Glu His Ile Leu Leu Ala Lys Gln Val Gly Val 115 120 125

Pro Asn Ile Val Val Phe Leu Asn Lys Glu Asp Met Val Asp Asp Ala 130 135 140

Glu Leu Leu Glu Leu Val Glu Leu Glu Val Arg Glu Leu Leu Ser Ser 145 150 155 160

Tyr Asp Phe Pro Gly Asp Asp Ile Pro Ile Val Ala Gly Ser Ala Leu 165 170 175

- Gln Ala Leu Glu Ala Ile Gln Gly Gly Ala Ser Gly Gln Lys Gly Asp 180 185 190
- Asn Pro Trp Val Asp Lys Ile Leu Lys Leu Met Glu Glu Val Asp Ala 195 200 205
- Tyr Ile Pro Thr Pro Glu Arg Glu Val Asp Arg Pro Phe Leu Met Ala 210 215 220
- Val Glu Asp Val Phe Thr Ile Thr Gly Arg Gly Thr Val Ala Thr Gly 225 230 235 240
- Arg Ile Glu Arg Gly Ser Val Lys Val Gly Glu Thr Ile Glu Ile Val 245 250 255
- Gly Leu Arg Asp Thr Arg Ser Thr Thr Val Thr Gly Val Glu Met Phe 260 265 270
- Gln Lys Thr Leu Asp Glu Gly Leu Ala Gly Asp Asn Val Gly Leu Leu 275 280 285
- Leu Arg Gly Ile Gln Lys Thr Asp Ile Glu Arg Gly Met Val Leu Ala 290 295 300
- Lys Pro Gly Ser Ile Thr Pro His Thr Lys Phe Glu Ser Glu Val Tyr 305 310 315 320
- Val Leu Lys Lys Glu Glu Gly Gly Arg His Thr Pro Phe Phe Pro Gly 325 330 335
- Tyr Arg Pro Gln Phe Tyr Val Arg Thr Thr Asp Val Thr Gly Ala Ile 340 345 350
- Ser Asp Phe Thr Ala Asp Asp Gly Ser Ala Ala Glu Met Val Ile Pro 355 360 365
- Gly Asp Arg Ile Lys Met Thr Val Glu Leu Ile Asn Pro Ile Ala Ile 370 375 380
- Glu Gln Gly Met Arg Phe Ala Ile Arg Glu Gly Gly Arg Thr Ile Gly 385 390 395 400
- Ala Gly Val Val Ser Lys Ile Leu Gln

<210> 4

<211> 408

<212> PRT

<213> Cryptomonas phi

<400> 4

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Ser Ala Thr Leu Ser Gln Tyr Thr Gly Lys Ser Lys Lys Phe Asp Glu 35 40 45

Ile Asp Ser Ala Pro Glu Glu Lys Ala Arg Gly Ile Thr Ile Asn Thr
50 55 60

Ala His Val Glu Tyr Glu Thr Asp Lys Trp Tyr Tyr Ala His Val Asp 65 70 75 80

Cys Pro Gly His Ala Asp Tyr Val Lys Asn Met Ile Thr Gly Ala Ala 85 90 95

Gln Met Asp Gly Ala Ile Leu Val Cys Ser Ala Ala Asn Gly Pro Met 100 105 110

Pro Gln Thr Arg Glu His Ile Leu Leu Ala Lys Gln Val Gly Val Pro 115 120 125

Tyr Ile Val Val Phe Leu Asn Lys Ala Asp Met Val Asp Asp Glu Glu 130 135 140

Leu Leu Glu Leu Val Gln Leu Glu Val Gln Glu Leu Leu Glu Lys Tyr 145 150 155 160

Asp Phe Pro Gly Ser Glu Ile Pro Phe Val Ala Gly Ser Ala Leu Leu 165 170 175

- Ala Leu Glu Ala Val Ala Asn Asn Pro Thr Ile Lys Arg Gly Glu Asp 180 185 190
- Lys Trp Val Asp Thr Ile Tyr Gln Leu Met Asp Lys Val Asp Glu Tyr 195 200 205
- Ile Pro Thr Pro Glu Arg Glu Thr Asp Lys Ala Phe Leu Met Ala Val 210 215 220
- Glu Asp Val Phe Ser Ile Thr Gly Arg Gly Thr Val Ala Thr Gly Arg 225 230 235 240
- Ile Glu Arg Gly Lys Val Lys Val Gly Asp Thr Ile Glu Ile Val Gly 245 250 255
- Leu Arg Glu Thr Arg Asn Thr Thr Ile Thr Gly Leu Glu Met Phe Gln 260 265 270
- Lys Ser Leu Asp Glu Ala Leu Ala Gly Asp Asn Val Gly Ile Leu Val 275 280 285
- Arg Gly Ile Gln Lys Thr Asp Ile Glu Arg Gly Met Val Leu Ala Ala 290 295 300
- Pro Gly Ser Ile Thr Pro His Thr Lys Phe Glu Gly Glu Val Tyr Val 305 310 315 320
- Leu Thr Lys Glu Glu Gly Gly Arg His Thr Pro Phe Phe Ser Gly Tyr 325 330 335
- Arg Pro Gln Phe Tyr Val Arg Thr Thr Asp Val Thr Gly Thr Ile Ala 340 345 350
- Gln Phe Thr Ser Asp Asp Gly Ser Thr Ala Glu Met Val Met Pro Gly 355 360 365
- Asp Arg Ile Lys Met Thr Ala Gln Leu Ile His Pro Ile Ala Ile Glu 370 375 380
- Lys Gly Met Arg Phe Ala Ile Arg Glu Gly Gly Arg Thr Val Gly Ala 385 390 395 400
- Gly Val Val Ser Lys Ile Ile Glu

<210> 5

<211> 409

<212> PRT

<213> Cyanophora paradoxa

<400> 5

Met Ala Arg Gln Lys Phe Asp Gly Asn Lys Pro His Val Asn Ile Gly
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Thr Ile Gly His Val Asp His Gly Lys Thr Thr Leu Thr Ala Ala Ile 20 25 30

Thr Thr Ala Leu Ala Ser Gln Gly Lys Gly Lys Ala Arg Lys Tyr Asp 35 40 45

Glu Ile Asp Ala Ala Pro Glu Glu Lys Ala Arg Gly Ile Thr Ile Asn 50 55 60

Thr Ala His Val Glu Tyr Glu Thr Glu Lys Arg His Tyr Ala His Val 65 70 75 80

Asp Cys Pro Gly His Ala Asp Tyr Val Lys Asn Met Ile Thr Gly Ala 85 90 95

Ala Gln Met Asp Gly Ala Ile Leu Val Val Ser Ala Ala Asp Gly Pro 100 105 110

Met Pro Gln Thr Arg Glu His Ile Leu Leu Ala Lys Gln Val Gly Val
115 120 125

Pro Asn Met Val Val Phe Leu Asn Lys Glu Asp Gln Ile Asp Asp Ala 130 135 140

Asp Leu Leu Glu Leu Val Glu Leu Glu Val Arg Glu Leu Leu Ser Lys 145 150 155 160

Tyr Asp Phe Pro Gly Asp Gln Ile Pro Phe Val Ser Gly Ser Ala Leu 165 170 175

- Leu Ala Leu Glu Ser Leu Ser Ser Asn Pro Lys Leu Met Arg Gly Glu 180 185 190
- Asp Lys Trp Val Asp Lys Ile Leu Ala Leu Met Asp Ala Val Asp Glu 195 200 205
- Tyr Ile Pro Thr Pro Glu Arg Pro Ile Asp Lys Ser Phe Leu Met Ala 210 215 220
- Ile Glu Asp Val Phe Ser Ile Thr Gly Arg Gly Thr Val Ala Thr Gly 225 230 235 240
- Arg Ile Glu Arg Gly Ala Ile Lys Val Gly Glu Thr Val Glu Leu Val 245 250 255
- Gly Leu Lys Asp Thr Lys Ser Thr Thr Val Thr Gly Leu Glu Met Phe 260 265 270
- Gln Lys Thr Leu Glu Glu Gly Met Ala Gly Asp Asn Ile Gly Ile Leu 275 280 285
- Leu Arg Gly Val Gln Lys Thr Asp Ile Glu Arg Gly Met Val Leu Ala 290 295 300
- Lys Pro Gly Ser Ile Thr Pro His Thr Gln Phe Glu Ser Glu Val Tyr 305 310 315 320
- Val Leu Thr Lys Asp Glu Gly Gly Arg His Thr Pro Phe Phe Ser Gly 325 330 335
- Tyr Arg Pro Gln Phe Tyr Val Arg Thr Thr Asp Val Thr Gly Ser Ile 340 345 350
- Asp Ala Phe Thr Ala Asp Asp Gly Ser Asn Ala Glu Met Val Met Pro 355 360 365
- Gly Asp Arg Ile Lys Met Thr Val Ser Leu Val His Pro Ile Ala Ile 370 375 380
- Glu Gln Gly Met Arg Phe Arg Ile Arg Glu Gly Gly Arg Thr Ile Gly 385 390 395 400
- Ala Gly Val Val Ser Lys Ile Leu Lys

<210> 6 <211> 394 <212> PRT <213> Escherichia coli

Thr Ile Gly His Val Asp His Gly Lys Thr Thr Leu Thr Ala Ala Ile 20 25 30

Thr Thr Val Leu Ala Lys Thr Tyr Gly Gly Ala Ala Arg Ala Phe Asp 35 40 45

Gln Ile Asp Asn Ala Pro Glu Glu Lys Ala Arg Gly Ile Thr Ile Asn 50 55 60

Thr Ser His Val Glu Tyr Asp Thr Pro Thr Arg His Tyr Ala His Val 65 70 75 80

Asp Cys Pro Gly His Ala Asp Tyr Val Lys Asn Met Ile Thr Gly Ala 85 90 95

Ala Gln Met Asp Gly Ala Ile Leu Val Val Ala Ala Thr Asp Gly Pro 100 105 110

Met Pro Gln Thr Arg Glu His Ile Leu Leu Gly Arg Gln Val Gly Val 115 120 125

Pro Tyr Ile Ile Val Phe Leu Asn Lys Cys Asp Met Val Asp Asp Glu 130 135 140

Glu Leu Leu Glu Leu Val Glu Met Glu Val Arg Glu Leu Leu Ser Gln 145 150 155 160

Tyr Asp Phe Pro Gly Asp Asp Thr Pro Ile Val Arg Gly Ser Ala Leu 165 170 175

- Lys Ala Leu Glu Gly Asp Ala Glu Trp Glu Ala Lys Ile Leu Glu Leu 180 185 190
- Ala Gly Phe Leu Asp Ser Tyr Ile Pro Glu Pro Glu Arg Ala Ile Asp 195 200 205
- Lys Pro Phe Leu Leu Pro Ile Glu Asp Val Phe Ser Ile Ser Gly Arg 210 215 220
- Gly Thr Val Val Thr Gly Arg Val Glu Arg Gly Ile Ile Lys Val Gly 225 230 235 240
- Glu Glu Val Glu Ile Val Gly Ile Lys Glu Thr Gln Lys Ser Thr Cys 245 250 255
- Thr Gly Val Glu Met Phe Arg Lys Leu Leu Asp Glu Gly Arg Ala Gly 260 265 270
- Glu Asn Val Gly Val Leu Leu Arg Gly Ile Lys Arg Glu Glu Ile Glu 275 280 285
- Arg Gly Gln Val Leu Ala Lys Pro Gly Thr Ile Lys Pro His Thr Lys 290 295 300
- Phe Glu Ser Glu Val Tyr Ile Leu Ser Lys Asp Glu Gly Gly Arg His 305 310 315 320
- Thr Pro Phe Phe Lys Gly Tyr Arg Pro Gln Phe Tyr Phe Arg Thr Thr 325 330 335
- Asp Val Thr Gly Thr Ile Glu Leu Pro Glu Gly Val Glu Met Val Met 340 345 350
- Pro Gly Asp Asn Ile Lys Met Val Val Thr Leu Ile His Pro Ile Ala 355 360 365
- Met Asp Asp Gly Leu Arg Phe Ala Ile Arg Glu Gly Gly Arg Thr Val 370 375 380
- Gly Ala Gly Val Val Ala Lys Val Leu Ser 385 390

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<211> 58
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<400> 9
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<211> 58
<212> RNA
<213> Plasmodium falciparum
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<211> 58
<212> RNA
<213> Toxoplasma gondii
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Asp Cys Pro Gly
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<213> Escherichia coli
<400> 14
Asn Lys Cys Asp
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